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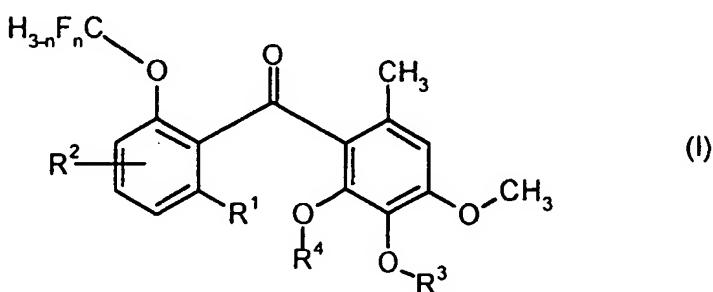
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(54) Fungicidal 2-methoxybenzophenones

(57) Substituted benzophenone compounds are disclosed, having the formula:



The compounds are useful as fungicides having high systemicities.

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Description

[0001] This invention relates to certain benzophenone compounds, a process for their preparation, compositions containing such compounds, a method for combating a fungus at a locus comprising treating the locus with such compounds and their use as fungicides.

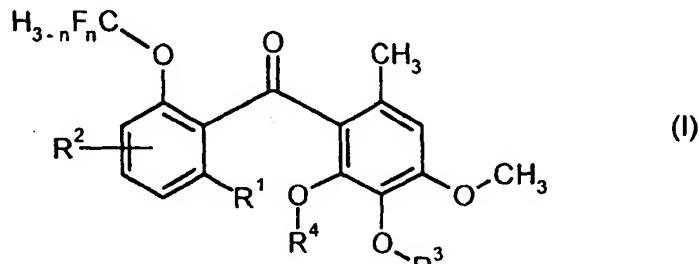
[0002] Food production relies upon a variety of agricultural technologies to ensure the growing population's dietary needs remain affordable, nutritious and readily available on grocery store shelves. Fungicides are one of these agricultural technologies which are available to the world community. Fungicides are agrochemical compounds which protect crops and foods from fungus and fungal diseases. Crops and food are constantly threatened by a variety of fungal organisms, which, if left uncontrolled, can cause ruined crops and devastated harvests.

[0003] In particular, ascomycetes, the causative agent for powdery mildew diseases are an ever-present threat especially to cereal and fruit crops. However, applications of fungicidal agents at disease control rates can cause phytotoxic damage to the target plants.

[0004] The compounds of the present invention are disclosed in a general formula of European patent ("EP") application EP 0 727 141, which published on August 21, 1996. The EP application discloses compounds having activity against phytopathogenic fungi, but relatively low systemicity.

[0005] There is no disclosure in the EP application of substituted benzophenones, wherein the first phenyl group is substituted by a methoxy group in the 2-position and by a halogen atom or an alkyl group in the 6-position and the second phenyl group is substituted by three alkoxy groups and one methyl group.

[0006] The present invention provides a compound of formula I



wherein

R^1 represents a halogen atom or an alkyl or haloalkyl group;

R^2 represents a hydrogen or halogen atom, or an alkyl, haloalkyl, alkoxy or nitro group; or R^1 and R^2 together represent a group of formula $-CH=CH-CH=CH-$:

R³ and R⁴ each independently represent an optionally substituted alkyl group; and

IV and V. Each independent

The compounds combine relatively excellent selective fungicidal activities in various crops with comparably high systemicities.

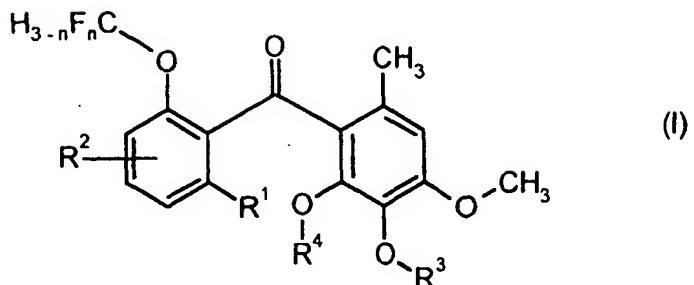
[0007] It is an object of the present invention to provide highly systemic fungicidal compounds.

[0008] It is also an object of the invention to provide methods for controlling an undesired fungus by contacting said plants with a fungicidally effective amount of the compounds.

[0009] It is another object of the invention to provide selective fungicidal compositions containing the compounds as active ingredients.

[0010] These and other objects and features of the invention will be more apparent from the detailed description set forth hereinbelow, and from the appended claims.

[0011] It has surprisingly been found that the compounds of formula I



in which R¹ through R⁴ and n have the meaning given above combine relatively excellent fungicidal activity against phytopathogenic fungi, in particular those that cause powdery mildew diseases and have comparably high systemicity.

[0012] In general terms, unless otherwise stated, as used herein the term halogen atom may denote a bromine, iodine, chlorine or fluorine atom, and is especially a bromine, chlorine or fluorine atom, in particular a bromine or chlorine atom.

[0013] Optionally substituted moieties may be unsubstituted or have from one up to the maximal possible number of substituents. Typically, 0 to 2 substituents are present. Each optionally substituted group independently is substituted by one or more halogen atoms or nitro, cyano, cycloalkyl, preferably C₃₋₆ cycloalkyl, cycloalkenyl, preferably C₃₋₆ cycloalkenyl, haloalkyl, preferably C₁₋₆ haloalkyl, halocycloalkyl, preferably C₃₋₆ halocycloalkyl, alkoxy, preferably C₁₋₆ alkoxy, haloalkoxy, preferably C₁₋₆ haloalkoxy, phenyl, halo- or dihalo-phenyl or pyridyl groups.

[0014] In general terms, unless otherwise stated herein, the terms alkyl and alkoxy as used herein with respect to a radical or moiety refer to a straight or branched chain radical or moiety. As a rule, such radicals have up to 10, in particular up to 6 carbon atoms. Suitably an alkyl or alkoxy moiety has from 1 to 6 carbon atoms, preferably from 1 to 5 carbon atoms. A preferred alkyl moiety is the methyl, ethyl, n-propyl, isopropyl or n-butyl group.

[0015] In general terms, unless otherwise stated herein, the term haloalkyl as used herein with respect to a radical or moiety refers to a straight or branched chain radical or moiety being substituted by one or more halogen atoms. As a rule, such radicals have up to 10, in particular up to 6 carbon atoms. Suitably haloalkyl moieties are fluoroalkyl moieties having from 1 to 6 carbon atoms, preferably from 1 or 2 carbon atoms. The fluoroalkyl groups include alkyl groups in which one or more, in particular all hydrogen atoms have been replaced by fluorine. A preferred fluoroalkyl moiety is represented by the formula $-(\text{CH}_2)_m-(\text{CF}_2)_n-\text{Y}$, in which m is 0, 1 or 2, n is 1, 2, 3, 4 or 5 and Y represents a hydrogen or fluoro atom. Particularly preferred is the difluoromethyl, trifluoromethyl, 2,2,2-trifluoroethyl, pentafluoroethyl, or *n*-heptafluoropropyl, group.

[0016] The invention especially relates to compounds of the general formula I in which any alkyl part of the groups R¹ through R⁴, which may be straight chained or branched, contains up to 10 carbon atoms, preferably up to 9 carbon atoms, more preferably up to 6 carbon atoms, and in which each optionally substituted group independently is substituted by one or more halogen atoms or nitro, cyano, cycloalkyl, preferably C₃₋₆ cycloalkyl, cycloalkenyl, preferably C₃₋₆ cycloalkenyl, haloalkyl, preferably C₁₋₆ haloalkyl, halocycloalkyl, preferably C₃₋₆ halocycloalkyl, alkoxy, preferably C₁₋₆ alkoxy, haloalkoxy, preferably C₁₋₆ haloalkoxy, phenyl, or pyridyl groups, in which the phenyl moiety is optionally substituted by one or two substituents selected from halogen atoms, cyano, C₁₋₆ alkyl and C₁₋₆ alkoxy groups.

[0017] The invention especially relates to compounds of the general formula I in which R¹ represents a halogen atom, in particular chlorine, a straight-chained or branched C₁₋₁₀ alkyl, in particular a straight-chained C₁₋₃ alkyl group, most preferably being a methyl group being unsubstituted or substituted by at least one optionally substituted phenyl group; or a straight-chained or branched C₁₋₁₀ fluoroalkyl, in particular a straight-chained C₁₋₃ fluoroalkyl group, most preferably being a trifluoromethyl group.

[0018] The invention especially relates to compounds of the general formula I in which R² represents a hydrogen or halogen atom, in particular a chlorine, bromine or iodine atom, a nitro, a C₁₋₁₀ alkyl or a C₁₋₁₀ haloalkyl group, most preferred being a hydrogen, chlorine or bromine atom or a trifluoromethyl group. If R² is different from hydrogen, it is most preferred attached in the ortho-position to radical R¹.

[0019] The invention especially relates to compounds of the general formula I in which R³ and R⁴ each independently represent an optionally substituted straight-chained or branched C₁₋₅ alkyl, in particular a straight-chained C₁₋₃ alkyl group, most preferably an unsubstituted or substituted methyl group. This methyl group is preferably unsubstituted or substituted by a phenyl group which is unsubstituted or substituted by one to five, preferably one or two halogen atoms or C₁₋₄ alkyl or C₁₋₄ alkoxy groups.

[0020] The benzophenone compounds according to formula I are oils, gums, or crystalline solid materials and possess valuable fungicidal properties. For example, they can be used in agriculture, or related fields such as horticulture and viticulture, for the control of phytopathogenic fungi, especially ascomycetes, in particular powdery mildew diseases.

such as those caused by *Blumeria (Erysiphe) graminis*, *Erysiphe cichoracearum*, *Podosphaera leucotricha*, *Uncinula necator* and the like. Said benzophenone compounds possess a high fungicidal activity over a wide concentration range and may be used in agriculture without harmful phytotoxic effects.

[0021] Moreover, the compounds according to the invention show enhanced curative and residual control of fungi and fungal diseases such as cereal, cucumber, apple and grape powdery mildew, and improved foliar systemicity compared with conventional fungicides.

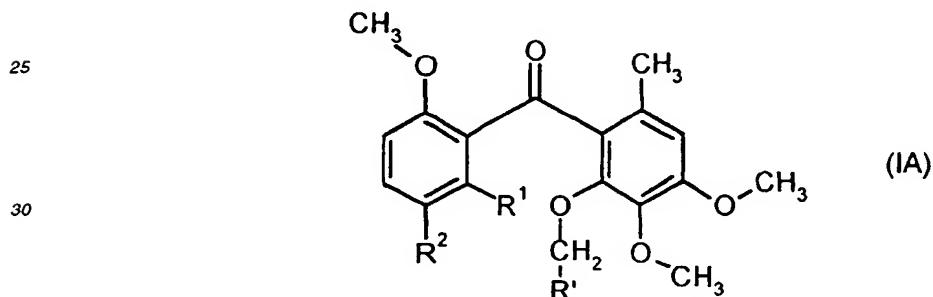
[0022] Useful results in terms of control of phytopathogenic fungi are obtained with a compound as defined in formula I wherein:

10 R¹ represents a chloro atom or a methyl or trifluoromethyl group;
 R² represents a hydrogen, chloro or bromo atom or trifluoromethyl group;
 R³ represents a C₁₋₅ alkyl group;
 R⁴ represents a C₁₋₅ alkyl group or a benzyl group being optionally substituted by one or more halogen atoms or one or more C₁₋₄ alkyl or alkoxy groups; and
 15 n is 0 or 2, in particular 0.

[0023] If R² represents Cl, Br or CF₃, it is preferably attached to the benzene ring in the ortho-position with respect to radical R¹.

[0024] Furthermore preferred are those compounds of formula I, wherein at least one of the radicals R¹ and R² represents a trifluoromethyl group

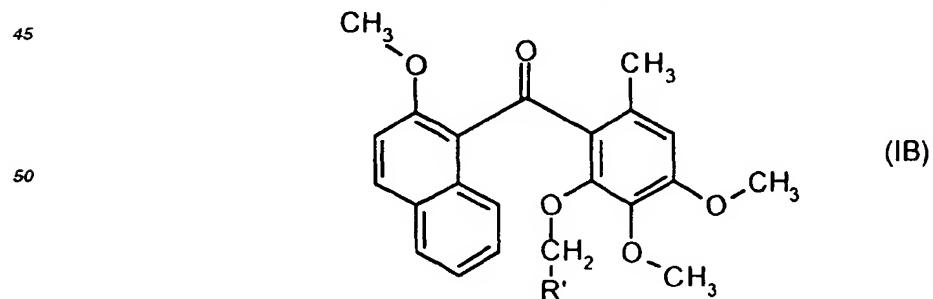
[0025] In particular the compounds of formula IA are preferred:



35 in which

R¹ represents a chloro atom or a methyl or trifluoromethyl group,
 R² represents a hydrogen, chloro or bromo atom or a methyl or trifluoromethyl group, and
 40 R' represents a hydrogen atom or a C₁₋₄ alkyl or a phenyl group being optionally substituted by one or more fluorine atoms or one or more C₁₋₄ alkyl groups.

[0026] In particular the compounds of formula IB are preferred:



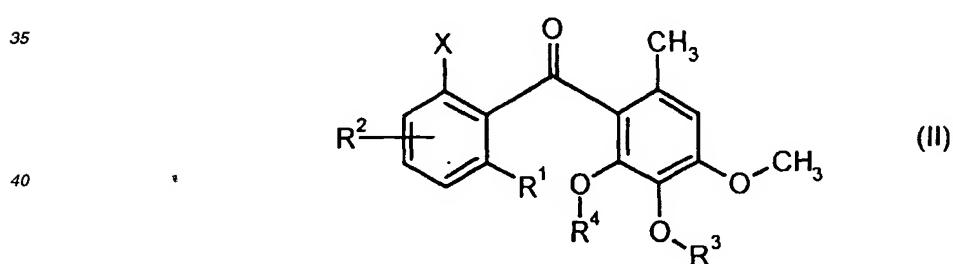
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in which R' represents a hydrogen atom or a C₁₋₄ alkyl or a phenyl group being optionally substituted by one or more fluorine atoms or one or more C₁₋₄ alkyl groups.

[0027] Especially good results in terms of control of phytopathogenic fungi are obtained by using, for example, the following compounds of formula I:

5 6,6'-dimethyl-2,2',3',4'-tetramethoxy-benzophenone,
 6,6'-dimethyl-3'-pentoxy-2,2',4'-trimethoxy-benzophenone,
 5-bromo-6,6'-dimethyl-2,2',3',4'-tetramethoxy-benzophenone,
 5-chloro-6,6'-dimethyl-2,2',3',4'-tetramethoxy-benzophenone,
 5-iodo-6,6'-dimethyl-2,2',3',4'-tetramethoxy-benzophenone,
 6-chloro-6'-methyl-2,2',3',4'-tetramethoxy-benzophenone,
 10 5-bromo-6-chloro-6'-methyl-2,2',3',4'-tetramethoxy-benzophenone,
 6-chloro-5,6'-dimethyl-2,2',3',4'-tetramethoxy-benzophenone,
 2'-n-butoxy-6-chloro-6'-methyl-2,3',4'-trimethoxy-benzophenone,
 2'-n-butoxy-6-chloro-5,6'-dimethyl-2,2',benzophenone
 6-chloro-2'-(2-fluorobenzoyloxy)-6'-methyl-2,3',4'-trimethoxy-benzophenone,
 15 6-chloro-2'-(4-fluorobenzoyloxy)-6'-methyl-2,3',4'-trimethoxy-benzophenone,
 5-bromo-6,6'-dimethyl-3'-n-pentyloxy-2,2',4'-trimethoxy-benzophenone
 6-chloro-6'-methyl-2'-n-pentyloxy-2,3',4'-trimethoxy-benzophenone,
 6-chloro-2'-(3-methylbutyloxy)-6'-methyl-2,3',4'-trimethoxy-benzophenone,
 2'-benzyloxy-6-chloro-6'-methyl-2,3',4'-trimethoxy-benzophenone,
 20 6-chloro-2'-(3-methylbenzyloxy)-6'-methyl-2,3',4'-trimethoxy-benzophenone,
 6-chloro-2'-(4-methylbenzyloxy)-6'-methyl-2,3',4'-trimethoxy-benzophenone,
 6-chloro-2-difluoromethoxy-6'-methyl-2,3',4'-trimethoxy-benzophenone,
 1-(6-methyl-2,3,4-trimethoxybenzoyl)-2-methoxynaphthalene
 1-(6-methyl-2,3,4-trimethoxybenzoyl)-2-difluoromethoxynaphthalene,
 25 5-chloro-6,6'-dimethyl-3'-ethoxy-2,2',4'-trimethoxy-benzophenone,
 5-chloro-6,6'-dimethyl-3'-n-propyloxy-2,2',4'-trimethoxy-benzophenone,
 6,6'-dimethyl-2,2',3',4'-tetramethoxy-5-trifluoromethyl-benzophenone,
 6'-methyl-2,2',3',4'-tetramethoxy-6-trifluoromethyl-benzophenone,
 5-bromo-6'-methyl-2,2',3',4'-tetramethoxy-6-trifluoromethylbenzophenone.

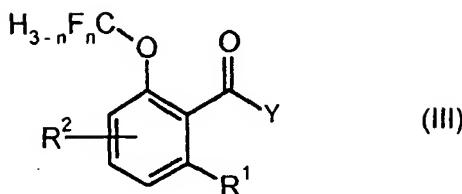
30 [0028] The present invention further provides a process (A) for the preparation of a compound of formula I, wherein n is 0, which comprises treating a compound of the general formula II



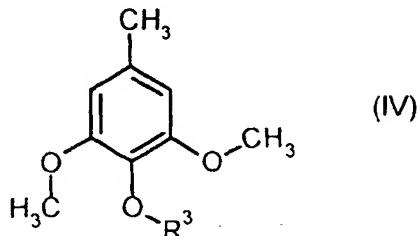
wherein R¹, R², R³ and R⁴ have the meaning given above and X represents a fluoro or chloro atom, with an alkali methylate, preferably sodium methylate.

[0029] Another possibility for the preparation of the compounds of formula I is a process (B) which comprises the steps of

(a) reacting a compound of formula III,

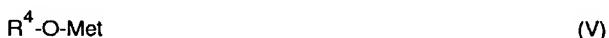


10 wherein R¹, R² and n have the meaning given above and Y represents a leaving group, in particular a chloro atom or a hydroxy group, with compound of formula IV,



wherein R³ has the meaning given for claim 1; in the presence of a Lewis acid (Y= leaving group) or a dehydrating agent (Y = OH), preferably phosphorous pentoxide or POCl₃; and

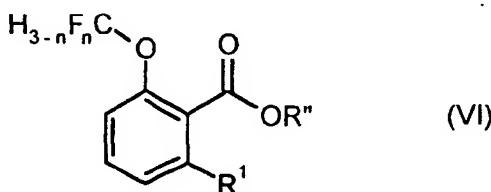
25 (b) optionally treating the resulting benzophenone of formula I, wherein R⁴ represents a methyl group, with a compound of formula V,



wherein R⁴ represents an optionally substituted alkyl group, being different from methyl, and Met denotes an alkali metal atom, preferably sodium.

35 [0030] The compounds of formula III, wherein R² represents a halogen atom, are preferably obtained by a process (C) which comprises the steps of

40 (a) reacting a compound of formula VI,



50 wherein R¹ and n have the meaning given for claim 1 and R'' represents a hydrogen atom or an alkyl group, with a halogenating agent,

55 (b) optionally hydrolyzing the resulting halogenated alkyl benzoate (R'' = alkyl), and
(c) optionally treating the resulting halogenated benzoic acid with thionyl chloride.

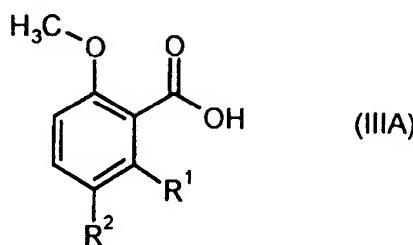
[0031] The compounds of formula I, wherein R² represents a trifluoromethyl group are preferably obtained by treating a compound of formula I, wherein R² represents a halogen atom, with trifluoroacetate in the presence of a copper salt at elevated temperatures.

[0032] The starting materials of formulae II, IV, V and VI are known products, the starting materials of formula III are partly known and partly novel products.

[0033] Accordingly, the invention provides novel intermediate of formula IIIA

5

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wherein R¹ represents an alkyl group, in particular a methyl group and R² represents a bromo atom or a trifluoromethyl group.

[0034] The starting materials of formulae II, IV, V and VI may be prepared according to established methods or routine adaptations thereof. Substituents which are not compatible with the selected reaction conditions may be introduced after formation of the benzophenone. They may be generated by known methods such as subsequent derivatization or substitution of a suitable group or by cleaving off a suitable protecting group.

[0035] The reaction between the 2-halobenzophenone of formula II and the alkali metal methylate is preferably carried out in the presence of a solvent, such as ethers like tetrahydrofuran, diethylether, *tert*-butyl-methylether or dimethoxyethane or methanol or in mixtures of these solvents. The molar ratio between formula II and the alkali metal methylate is preferably in the range of 0.3 to 1.9 at a temperature between 25 and 120 °C.

[0036] The Friedel Crafts reaction between formula III and IV is effected in the presence of a Lewis acid catalyst according to well-established procedures (Y = Cl). Suitable catalysts include FeCl₃, AlCl₃, SnCl₄, ZnCl₂, TiCl₄, SbCl₅ and BF₃, which may be in a molar equivalent amount (based on the benzoyl chloride of formula III). However, it is also possible to use lesser amounts of catalyst at elevated temperatures, suitably under reflux temperatures, preferred catalysts under these conditions being FeCl₃, I₂, ZnCl₂, iron, copper, strong sulphonic acids such as F₃CSO₃H, and acidic ion exchange resins such as Amberlyst®15 and Nafion®. The preferred catalyst is FeCl₃ in a 0.001 to 0.2 molar ratio at a temperature of about 50 to 180°C. The reaction can be carried out in a solvent inert under the reaction conditions, for example ethylene or methylene chloride, benzene, octane, decane or solvent mixtures, or in the absence of solvent, conveniently by employing one of the reactants in excess, e. g. in the range of 1:5 to 5:1. If AlCl₃ is being used, the molar ratio is preferably in the range of 0.5 to 2 and the suitable solvents are e.g. dichloromethane or ethylenechloride at a temperature usually between -10 and 70°C.

[0037] In another preferred process according to the invention the benzoic acid of formula III (Y = OH) is reacted with a compound of formula IV in the presence of phosphorous pentoxide at temperatures of about 0 to 50 °C, preferably at room temperature or in the presence of POCl₃ at temperatures of about 50 to 150 °C, preferably under reflux.

[0038] The halogenation of the benzoate of formula VI is preferably carried out in the presence of an inert solvent. Preferred halogenating agents are for example sulfonyl chloride, bromine and N-halo-succinimide. If R¹ represents a halogen atom highly polar solvents such as alcohols or carboxylic acids, in particular acetic acid are preferred. If R¹ represents an alkyl group, in particular a methyl group, apolar solvents such as tetrachloromethane are preferred. If the reaction is carried out with bromine at a temperature between 0 and 40 °C, preferably at room temperature, the halogenation takes place predominately in the ortho-position with respect to radical R¹.

[0039] In a preferred embodiment the compounds of formula III, in which R² represents a bromo atom are prepared by reacting a compound of formula VI, wherein R¹ represents an alkyl group, n is 0, and Rⁿ represents a hydrogen atom, with bromine. This bromination step is preferably carried out in the presence of a polar, protic solvent, such as an aliphatic alcohol or an aliphatic carboxylic acid, in particular acetic acid. The bromination may be carried out advantageously in the presence of a weak base or a buffer system, such as sodium acetate or sodium carbonate.

[0040] One embodiment of the process of the instant invention is a process wherein bromine is employed in an amount selected from a value in the range between 1.0 to 1.5, in particular 1.05 to 1.2 molar equivalents with respect to starting compound of formula VI.

[0041] The reaction between the compound of formula VI and bromine is as a rule carried out at a temperature sufficient to optimally convert the compound of formula VI to the compound of formula III. This term represents a temperature sufficiently high to maintain the conversion, but also sufficiently low to avoid decomposition of the starting material and the product. The reaction is carried out preferably at temperatures between 0 °C and 40 °C, in particular at ambient temperature.

[0042] The reaction between the compound of formula VI and bromine is as a rule carried out at a length of time sufficient to optimally convert the compound of formula VI to the compound of formula III. This term represents a period

of time sufficiently long to convert the maximum amount of the starting material to the compound of formula III. The reaction is carried out preferably at reaction time between 1 and 40 hours, in particular between 5 and 24 hours.

[0043] The transformation of a compound of formula I, wherein R² represents a halogen atom, into a compound of formula I, wherein one of R² represents a trifluoromethyl group, is carried out analogously to J. Chem. Soc. Perkin

5 Trans. PT1 (1988) N4, page 924. Preferably sodium trifluoroacetate is reacted with the halogen compound in the presence of a copper salt, preferably copper-(I) halides, in particular copper-(I) iodide. As a rule the reaction is carried out in a high-boiling polar aprotic solvent such as 1-methyl-2-pyrrolidone. The reaction is carried out preferably at temperatures between 120 °C and 200 °C, in particular between 150 °C and 170 °C.

[0044] The reaction between the compound of formula I, R² represents a halogen atom, and sodium trifluoroacetate 10 is as a rule carried out at a length of time sufficient to optimally transform the halogen compound into a trifluoromethyl compound. This term represents a period of time sufficiently long to convert the maximum amount of the starting material to the compound of formula I, wherein R² represents a trifluoromethyl group. The reaction is carried out preferably at a reaction time between 40 and 140 hours, in particular between 60 and 100 hours.

[0045] The processes described above and below can analogously be applied to other starting compounds, if desired.

[0046] Due to excellent plant tolerance, the compounds of formula I may be used in cultivation of all plants where 15 infection by the controlled fungi is not desired, e.g. cereals, vegetables, legumes, apples, vine. The absence of target crop phytotoxicity at fungus control rates is a feature of the present invention.

[0047] Accordingly the invention further provides a fungicidal composition which comprises a carrier and, as active 20 ingredient, at least one compound of formula I as defined above. A method of making such a composition is also provided which comprises bringing a compound of formula I as defined above into association with at least one carrier. Such a composition may contain a single compound or a mixture of several compounds of the present invention. It is also envisaged that different isomers or mixtures of isomers may have different levels or spectra of activity and thus compositions may comprise individual isomers or mixtures of isomers.

[0048] A composition according to the invention preferably contains from 0.5% to 95% by weight (w/w) of active 25 ingredient.

[0049] A carrier in a composition according to the invention is any material with which the active ingredient is formulated to facilitate application to the locus to be treated, which may for example be a plant, seed or soil, or to facilitate storage, transport or handling. A carrier may be a solid or a liquid, including material which is normally a gas but which has been compressed to form a liquid.

[0050] The compositions may be manufactured into e.g. emulsion concentrates, solutions, oil in water emulsions, 30 wettable powders, soluble powders, suspension concentrates, dusts, granules, water dispersible granules, microcapsules, gels and other formulation types by well-established procedures. These procedures include intensive mixing and/or milling of the active ingredients with other substances, such as fillers, solvents, solid carriers, surface active compounds (surfactants), and optionally solid and/or liquid auxiliaries and/or adjuvants. The form of application such as spraying, atomizing, dispersing or pouring may be chosen like the compositions according to the desired objectives 35 and the given circumstances.

[0051] Solvents may be aromatic hydrocarbons, e.g. Solvesso® 200, substituted naphthalenes, phthalic acid esters, such as dibutyl phthalate, aliphatic hydrocarbons, e.g. cyclohexane or paraffins, alcohols and glycols as well 40 as their ethers and esters, e.g. ethanol, ethyleneglycol mono- and dimethyl ether, ketones such as cyclohexanone, strongly polar solvents such as N-methyl-2-pyrrolidone, or γ -butyrolactone, higher alkyl pyrrolidones, e.g. n-octylpyrrolidone or cyclohexylpyrrolidone, epoxidized plant oil esters, e.g. methylated coconut or soybean oil ester and water. Mixtures of different liquids are often suitable.

[0052] Solid carriers, which may be used for dusts, wettable powders, waterdispersible granules, or granules, may 45 be mineral fillers, such as calcite, talc, kaolin, montmorillonite or attapulgite. The physical properties may be improved by addition of highly dispersed silica gel or polymers. Carriers for granules may be porous material, e.g. pumice, kaolin, sepiolite, bentonite; non-sorptive carriers may be calcite or sand. Additionally, a multitude of pre-granulated inorganic or organic materials may be used, such as dolomite or crushed plant residues.

[0053] Pesticidal compositions are often formulated and transported in a concentrated form which is subsequently 50 diluted by the user before application. The presence of small amounts of a carrier which is a surfactant facilitates this process of dilution. Thus, preferably at least one carrier in a composition according to the invention is a surfactant. For example, the composition may contain at two or more carriers, at least one of which is a surfactant.

[0054] Surfactants may be nonionic, anionic, cationic or zwitterionic substances with good dispersing, emulsifying and wetting properties depending on the nature of the compound according to general formula I to be formulated. Surfactants may also mean mixtures of individual surfactants.

[0055] The compositions of the invention may for example be formulated as wettable powders, water dispersible 55 granules, dusts, granules, solutions, emulsifiable concentrates, emulsions, suspension concentrates and aerosols. Wettable powders usually contain 5 to 90% w/w of active ingredient and usually contain in addition to solid inert carrier, 3 to 10% w/w of dispersing and wetting agents and, where necessary, 0 to 10% w/w of stabilizer(s) and/or other additives

such as penetrants or stickers. Dusts are usually formulated as a dust concentrate having a similar composition to that of a wettable powder but without a dispersant, and may be diluted in the field with further solid carrier to give a composition usually containing 0.5 to 10% w/w of active ingredient. Water dispersible granules and granules are usually prepared to have a size between 0.15 mm and 2.0 mm and may be manufactured by a variety of techniques. Generally,

5 these types of granules will contain 0.5 to 90% w/w active ingredient and 0 to 20% w/w of additives such as stabilizer, surfactants, slow release modifiers and binding agents. The so-called "dry flowables" consist of relatively small granules having a relatively high concentration of active ingredient. Emulsifiable concentrates usually contain, in addition to a solvent or a mixture of solvents, 1 to 80% w/v active ingredient, 2 to 20% w/v emulsifiers and 0 to 20% w/v of other additives such as stabilizers, penetrants and corrosion inhibitors. Suspension concentrates are usually milled so as to obtain a stable, non-sedimenting flowable product and usually contain 5 to 75% w/v active ingredient, 0.5 to 15% w/v of dispersing agents, 0.1 to 10% w/v of suspending agents such as protective colloids and thixotropic agents, 0 to 10% w/v of other additives such as defoamers, corrosion inhibitors, stabilizers, penetrants and stickers, and water or an organic liquid in which the active ingredient is substantially insoluble; certain organic solids or inorganic salts may be present dissolved in the formulation to assist in preventing sedimentation and crystallization or as antifreeze agents for water.

10 [0056] Aqueous dispersions and emulsions, for example compositions obtained by diluting the formulated product according to the invention with water, also lie within the scope of the invention.

15 [0057] Of particular interest in enhancing the duration of the protective activity of the compounds of this invention is the use of a carrier which will provide slow release of the pesticidal compounds into the environment of a plant which is to be protected.

20 [0058] The biological activity of the active ingredient can also be increased by including an adjuvant in the spray dilution. An adjuvant is defined here as a substance which can increase the biological activity of an active ingredient but is not itself significantly biologically active. The adjuvant can either be included in the formulation as a coformulant or carrier, or can be added to the spray tank together with the formulation containing the active ingredient.

25 [0059] As a commodity the compositions may preferably be in a concentrated form whereas the end user generally employs diluted compositions. The compositions may be diluted to a concentration down to 0.001% of active ingredient. The doses usually are in the range from 0.01 to 10 kg a.i./ha.

[0060] Examples of formulations according to the invention are:

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Emulsion Concentrate (EC)			
5	Active Ingredient Emulsifier(s)	Compound of Example 6 Atlox® 4856 B / Atlox® 4858 B 1) (mixture containing calcium alkyl aryl sulfonate, fatty alcohol ethoxylates and light aromatics / mixture containing calcium alkyl aryl sulfonate, fatty alcohol ethoxylates and light aromatics)	30 % (w/v) 5 % (w/v)
10	Solvent	shellsol® A 2) (mixture of C ₉ - C ₁₀ aromatic hydrocarbons)	to 1000 ml
Suspension Concentrate (SC)			
15	Active Ingredient Dispersing agent	Compound of Example 6 Soprophor® FL 3) (polyoxyethylene polyaryl phenyl ether phosphate amine salt)	50 % (w/v) 3 % (w/v)
20	Antifoaming agent Structure agent	Rhodorsil® 422 3) (nonionic aqueous emulsion of polydimethylsiloxanes) Kelzan® S 4) (Xanthan gum)	0.2 % (w/v) 0.2 % (w/v)
25	Antifreezing agent Biocidal agent	Propylene glycol proxef® 5) (aqueous dipropylene glycol solution containing 20% 1,2-benisothiazolin-3-one)	5 % (w/v) 0.1 % (w/v)
30	Water		to 1000 ml
Wettable Powder (WP)			
35	Active Ingredient Wetting agent	Compound of Example 6 Atlox® 49951) (polyoxyethylene alkyl ether)	60 % (w/w) 2 % (w/w)
40	Dispersing agent	Witcosperse® D-60 6) (mixture of sodium salts of condensed naphthalene sulfonic acid and alkylarylpolyoxy acetates)	3 % (w/w)
45	Carrier / Filler	Kaolin	35 % (w/w)
Water Dispersible Granules (WG)			
50	Active Ingredient Dispersing / Binding agent	Compound of Example 6 Witcosperse® D-450 6) (mixture of sodium salts of condensed naphthalene sulfonic acid and alkyl sulfonates)	50 % (w/w) 8 % (w/w)
55	Wetting agent Antifoaming agent	Morwet® EFW 6) (formaldehyde condensation product) Rhodorsil® EP 6703 3) (encapsulated silicone)	2 % (w/w) 1 % (w/w)
	Disintegrant	Agrimer® ATF 7) (cross-linked homopolymer of N-vinyl-2-pyrrolidone)	2 % (w/w)
	Carrier / Filler	Kaolin	35 % (w/w)

1) available from ICI Surfactants

2) available from Deutsche Shell AG

3) available from Rhodia former (Rhone-Poulenc)

4) available from Kelco Co.

5) available from Zeneca

6) available from Witco

7) from International Speciality Products

[0061] The compositions of this invention can also comprise other compounds having biological activity, e.g. compounds having similar or complementary pesticidal activity or compounds having plant growth regulating, fungicidal or

insecticidal activity. These mixtures of pesticides can have a broader spectrum of activity than the compound of general formula I alone. Furthermore, the other pesticide can have a synergistic effect on the pesticidal activity of the compound of general formula I.

[0062] The other fungicidal compound can be, for example, one which is capable of combating diseases of cereals (e.g. wheat) such as those caused by *Erysipha*, *Puccinia*, *Septoria*, *Gibberella* and *Helminthosporium* spp., seed and soil borne diseases and downy and powdery mildews on vines and powdery mildew and scab on apples etc. These mixtures of fungicides can have a broader spectrum of activity than the compound of general formula I alone.

[0063] Examples of the other fungicidal compounds are anilazine, azoxystrobin, benalaxyl, benomyl, binapacryl, bitertanol, blasticidin S, Bordeaux mixture, bromuconazole, bupirimate, captafol, captan, carbendazim, carboxin, carpropamid, chlorbenzthiazon, chlorothalonil, chlozolinate, copper-containing compounds such as copper oxychloride, and copper sulfate, cycloheximide, cymoxanil, cyprofuram, cyproconazole, cyprodinil, dichlofluanid, dichlone, dichloran, diclobutrazol, diclocymet, diclomezine, diethofencarb, difenoconazole, diflumetorim, dimethirimol, dimethomorph, diniconazole, dinocap, ditalimfos, dithianon, dodemorph, dodine, edifenphos, epoxiconazole, etaconazole, ethirimol, etridiazole, famoxadone, fenapanil, fenarimol, fenbuconazole, fenfuram, fenhexamid, fenpiclonil, fenpropidin, fenpropimorph, fentin, fentin acetate, fentin hydroxide, ferimzone, fluazinam, fludioxonil, flumetover, fluquinconazole, flusilazole, flusulfamide, flutolanil, flutriafol, folpet, fosetyl-aluminium, fuberidazole, furalaxy, furametyl, guazatine, hexaconazole, imazalil, iminoctadine, ipconazole, iprodione, isoprothiolane, kasugamycin, kitazin P, kresoxim-methyl, mancozeb, maneb, mepanipyrim, mepronil, metalaxyl, metconazole, methfuroxam, myclobutanil, neoasozin, nickel dimethyl-dithiocarbamate, nitrothalisopropyl, nuarimol, ofurace, organo mercury compounds, oxadixyl, oxycarboxin, penconazole, pencycuron, phenazineoxide, phthalide, polyoxin D, polyram, probenazole, prochloraz, procymidone, propamocarb, propiconazole, propineb, pyrazophos, pyrifenoxy, pyrimethanil, pyroquilon, pyroxyfur, quinomethionate, quinoxifen, quintozeno, spiroxamine, SSF-126, SSF-129, streptomycin, sulfur, tebuconazole, tecloftalamine, tecnazene, tetraconazole, thiabendazole, thifluzamide, thiophanate-methyl, thiram, tolclofosmethyl, tolylfluanid, triadimefon, triadimenol, triazbutil, triazoxide, tricyclazole, tridemorph, triflumizole, triforine, triticonazole, validamycin A, vinclozolin, XRD-563, zarilamid, zineb, ziram.

[0064] In addition, the co-formulations according to the invention may contain at least one benzophenone of formula I and any of the following classes of biological control agents such as viruses, bacteria, nematodes, fungi, and other microorganism which are suitable to control insects, weeds or plant diseases or to induce host resistance in the plants. Examples of such biological control agents are: *Bacillus thuringiensis*, *Verticillium lecanii*, *Autographica californica NPV*, *Beauvaria bassiana*, *Ampelomyces quisqualis*, *Bacillus subtilis*, *Pseudomonas fluorescens*, *Steptomyces griseoviridis* and *Trichoderma harzianum*.

[0065] Moreover, the co-formulations according to the invention may contain at least one benzophenone of formula I and a chemical agent that induces the systemic acquired resistance in plants such as for example nicotinic acid or derivatives thereof, 2,2-dichloro-3,3-dimethylcyclopropylcarboxylic acid or BION.

[0066] The compounds of general formula I can be mixed with soil, peat or other rooting media for the protection of the plants against seed-borne, soil-borne or foliar fungal diseases.

[0067] The invention still further provides the use as a fungicide of a compound of the general formula I as defined above or a composition as defined above, and a method for combating fungus at a locus, which comprises treating the locus, which may be for example plants subject to or subjected to fungal attack, seeds of such plants or the medium in which such plants are growing or are to be grown, with such a compound or composition.

[0068] The present invention is of wide applicability in the protection of crop and ornamental plants against fungal attack. Typical crops which may be protected include vines, grain crops such as wheat and barley, rice, sugar beet, top fruit, peanuts, potatoes, vegetables and tomatoes. The duration of the protection is normally dependent on the individual compound selected, and also a variety of external factors, such as climate, whose impact is normally mitigated by the use of a suitable formulation.

[0069] The following examples further illustrate the present invention. It should be understood, however, that the invention is not limited solely to the particular examples given below.

Example 1

Preparation of 6,6'-dimethyl-2,2',3',4'-tetramethoxy-benzophenone 1A 2-Methoxy-6-methylbenzoic acid

[0070] A mixture of ethyl 2-methoxy-6-methylbenzoate (5.0 g, 25 mmol), water (10 ml), methanol (40 ml) and sodium hydroxide (2.1 g, 50 mmol) is heated under reflux with stirring. The reaction mixture is diluted with water (150 ml) and acidified with concentrated hydrochloric acid. The solid material is collected by filtration, washed with water and dried yielding dark yellow crystals, 2.1 g, mp 136 °C.

1B 2-Methoxy-6-methylbenzoyl chloride

[0071] A mixture of 1A (1.7 g, 10.2 mmol) and thionyl chloride (2 ml) is heated under reflux for 1 hour. The mixture is concentrated and the resulting benzoylchloride is used without further purification.

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1C 6,6'-dimethyl-2,2',3',4'-tetramethoxy-benzophenone

[0072] A mixture of 3,4,5-trimethoxytoluene (1.86 g; 10.2 mmol), 1B (10.2 mmol), aluminium chloride (1.33 g, 10 mmol) and dichloromethane (20 ml) is stirred at 0° C. The reaction sets up at 0 °C with formation of hydrogen chloride.

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Subsequently, the reaction mixture is stirred for another 4 hours at room temperature. A mixture of dilute hydrochloric acid and ethyl acetate (1:1 v/v; 100 ml) is then slowly added at 0°C. The organic phase is concentrated and the residue is recrystallized from methanol. The solid material is collected by vacuum filtration, three times washed with methanol/water (3:1 v/v; 100 ml each) and dried yielding white crystals, 1.0 g (30.3 %), mp 84 °C.

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Example 2Preparation of 6,6'-dimethyl-2'-n-butoxy-2,3',4'-trimethoxy-benzophenone

[0073] A mixture of n-butanol (5 ml) and sodium hydride (60 % in oil, 10 mmol) is stirred until the formation of H₂ gas ceases. A mixture of 1C (0.7 g, 2.2 mmol) and dimethoxyethane (15 ml) is added to the resulting reaction mixture. Subsequently, the reaction mixture is heated under reflux with stirring for 24 hours. A mixture of water and ethyl acetate (1:1 v/v; 100 ml) is then slowly added at room temperature. The organic phase is separated, concentrated and the residue is purified by column chromatography (petrol ether : ethyl acetate, 95:5 v/v) yielding the pure product as a yellow oil, 0.2 g, (24.4 %).

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Example 3Preparation of 6-chloro-2'-pentyloxy-6'-methyl-2,3',4'-trimethoxy-benzophenone

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[0074] A mixture of sodium methylate in methanol (5.4 mol/l, 19.6 mmol), 2,6-dichloro-3',4'-dimethoxy-6'-methyl-2'-pentyloxy-benzophenone (obtained according to EP 0 727 141, 2.69 g, 6.5 mmol) and dimethoxyethane (20 ml) is heated to 80 °C with stirring for 24 hours. A mixture of water and ethyl acetate (1:1 v/v; 100 ml) is then slowly added at room temperature. The organic phase is separated and concentrated and the residue is purified by column chromatography (dichloromethane) yielding the pure product as a yellow oil, 0.52 g, (19.7 %).

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Example 4Preparation of 5-bromo-6-chloro-6'-methyl-2,2',3',4'-tetramethoxy-benzophenone

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4A Ethyl 5-bromo-6-chloro-2-methoxybenzoate

[0075] A mixture of ethyl 6-chloro-2-methoxybenzoate (1.8 g, 8.4 mmol), bromine (1.41 g, 8.8 mmol) and acetic acid (5 ml) is stirred at room temperature for 24 hours. The reaction mixture is poured into water and extracted with ethyl acetate. The organic phase is separated and concentrated and the residue is purified by column chromatography (petrol ether : ethyl acetate, 95:5 v/v) yielding the pure product as a yellow oil, 1.7 g, (69 %).

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4B 5-Bromo-6-chloro-2-methoxybenzoic acid

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[0076] A mixture of 4A (1.7 g, 5.8 mmol), water (10 ml), ethanol (20 ml) and sodium hydroxide (0.5 g, 12.5 mmol) is heated under reflux with stirring. The reaction mixture is diluted with water (80 ml) and acidified with concentrated hydrochloric acid. The solid material is collected by filtration, washed with water and dried yielding white crystals, 1.3 g (85 %), mp 186-188 °C.

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4C 5-Bromo-6-chloro-2-methoxybenzoyl chloride

[0077] A mixture of 4B (1.2 g, 4.6 mmol), dichloromethane (15 ml) and oxalyl chloride (1 ml) is stirred at room temperature for 2 hours. The mixture is concentrated and the resulting benzoylchloride is used without further purification.

4D 5-bromo-6-chloro-6'-methyl-2,2'3',4'-tetramethoxy-benzophenone

[0078] A mixture of 3,4,5-trimethoxytoluene (0.83 g; 4.6 mmol), 4C (4.6 mmol), aluminium chloride (0.62 g, 4.6 mmol) and dichloromethane (20 ml) is stirred at room temperature for 3 hours. A mixture of water and ethyl acetate (1:1 v/v; 50 ml) is then added. The organic phase is concentrated and the residue is crystallized from diisopropylether and recrystallized from methanol. The solid material is collected by vacuum filtration, washed with water and dried yielding yellow crystals, 0.7 g, (35.4 %) mp 87-88 °C.

Example 5Preparation of 1-(6'-methyl-2',3',4'-trimethoxybenzoyl)-2-methoxynaphthalene5A Methyl (2-methoxynaphth-1-yl)-carboxylate

[0079] A mixture of 2-hydroxynaphth-1-ylcarboxylic acid (18.82 g, 100 mmol), sodium hydroxide (8.8 g, 220 mmol), dimethylsulfate (31.5 g, 250 mmol) and water (200 ml) is heated to 70 °C with stirring for 20 hours. Subsequently, the reaction mixture is cooled to room temperature and extracted with ethyl acetate twice. The combined organic phases are concentrated and the residue is used without further purification.

5B Methyl (2-methoxynaphth-1-yl)-carboxylic acid

[0080] A mixture of crude 5A (10.5 g, 48 mmol), water (100 ml), methanol (150 ml) and sodium hydroxide (12 g, 300 mmol) is heated under reflux with stirring. The reaction mixture is extracted with diethylether twice. The aqueous reaction mixture is filtered and acidified with concentrated hydrochloric acid. The solid material is collected by filtration, washed with water and dried yielding yellow crystals, 9.45 g (97.4 %), mp 175-176 °C.

5C 1-(6'-methyl-2',3',4'-trimethoxybenzoyl)-2-methoxynaphthalene

[0081] A mixture of 5B (2.02 g, 10 mmol), 3,4,5-trimethoxytoluene (1.82 g, 10 mmol), P_2O_5 (10.0 g) and dichloromethane is stirred at room temperature for 16 hours. Subsequently, the dichloromethane is distilled off and the residue is diluted with ethyl acetate. The organic phase is washed with water and concentrated. The residue is purified by column chromatography (petrol ethers:ethyl acetate, 8:2 v/v) and recrystallized from petrol ether: diisopropylether (1:1 v/v). The solid material is collected by vacuum filtration, washed with cold petrol ether: diisopropylether (1:1 v/v) and dried, yielding white crystals, 0.9 g, (24.6 %) mp 72 °C.

Example 6Preparation of 5-bromo-6,6'-dimethyl-2,2'3',4'-tetramethoxy-benzophenone6A Ethyl 5-bromo-6-methyl-2-methoxybenzoate

[0082] A mixture of ethyl 6-methyl-2-methoxybenzoate (8.4 g, 43.2 mmol), bromine (6.9 g, 43.2 mmol) and tetrachloromethane (170 ml) is stirred at room temperature for 60 hours. The reaction mixture is poured into water and extracted with ethyl acetate. The organic phase is separated and concentrated. The crude product is obtained as a yellow oil, 10.3 g (87 %) and is used without further purification.

6B 5-Bromo-6-methyl-2-methoxybenzoic acid

[0083] A mixture of 6A (9.8 g, 34.1 mmol), water (40 ml), ethanol (80 ml) and sodium hydroxide (2.7 g, 68.3 mmol) is heated under reflux with stirring for 42 hours. The reaction mixture is diluted with water (80 ml), acidified with concentrated hydrochloric acid and extracted with dichloromethane. The organic phase is separated and concentrated. The solid material is collected by filtration, washed with water and dried yielding off-white crystals, 5.4 g (61 %), mp 81-83°C.

6C 5-bromo-6,6'-dimethyl-2,2'3',4'-tetramethoxy-benzophenone

[0084] A mixture of 6B (24 g, 10 mmol), 3,4,5-trimethoxytoluene (1.82 g; 10 mmol), P_2O_5 (10.0 g) and dichloromethane (150 ml) is stirred at room temperature for 16 hours. Subsequently, the dichloromethane is distilled off and the

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residue is diluted with ethyl acetate. The organic phase is washed with water and concentrated. The residue is purified by column chromatography (petrol ether: ethyl acetate, 8:2 v/v) and recrystallized from diisopropylether. The solid material is collected by vacuum filtration, washed with cold petrol ethers:diisopropylether (1:1 v/v) and dried, yielding white crystals, 2.2 g (54 %), mp 89-91 °C.

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Examples 7-62

[0085] Using essentially the same procedures described hereinabove for Examples 1 to 6 and employing standard derivatization techniques where appropriate, the following compounds are prepared and shown in Tables I and II:

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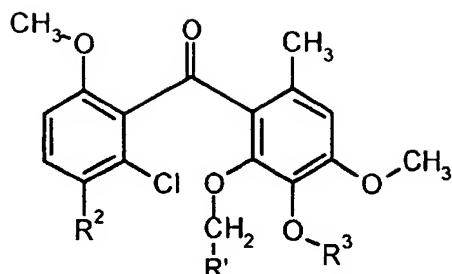
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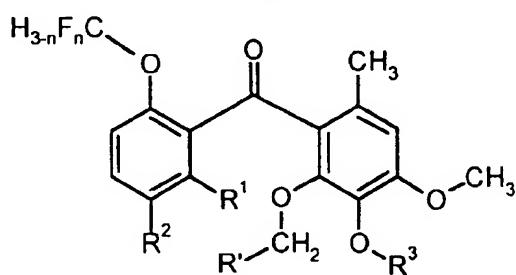
Table I



Example	R'	R ²	R ³	Melting point (° C)
7	H	methyl	methyl	95
8	n-propyl	methyl	methyl	oil
9	n-butyl	methyl	methyl	oil
10	H	H	methyl	51
11	n-propyl	H	methyl	oil
12	2-methylpropyl	H	methyl	55-56
13	phenyl	H	methyl	120-122
14	4-fluorophenyl	H	methyl	96-98
15	4-methylphenyl	H	methyl	80
16	3-methylphenyl	H	methyl	oil
17	2-fluorophenyl	H	methyl	90-93
18	2-methylpropyl	methyl	methyl	
19	phenyl	methyl	methyl	
20	4-fluorophenyl	methyl	methyl	
21	4-methylphenyl	methyl	methyl	
22	3-methylphenyl	methyl	methyl	
23	2-fluorophenyl	methyl	methyl	
24	n-propyl	Br	methyl	
25	n-butyl	Br	methyl	
26	2-methylpropyl	Br	methyl	
27	phenyl	Br	methyl	
28	4-fluorophenyl	Br	methyl	

29	4-methylphenyl	Br	methyl	
30	3-methylphenyl	Br	methyl	
31	2-fluorophenyl	Br	methyl	
32	n-propyl	H	n-butyl	oil
33	H	methyl	CO-ethyl	113-114
34	H	methyl	H	142-143
35	H	methyl	n-propyl	oil
36	H	methyl	n-butyl	oil
37	H	methyl	n-pentyl	oil
38	H	methyl	3-methyl- butyl	oil
39	H	NO ₂	methyl	oil
40	H	Cl	methyl	87
41	n-heptyl	H	methyl	oil
42	n-pentyl	H	methyl	oil

Table II



Example	n	R ¹	R ²	R'	R ³	Melting point (° C)
43	0	methyl	Br	H	n-pentyl	oil
44	0	methyl	isopropyl	H	methyl	
45	2	Cl	H	H	methyl	oil
46	2	-CH=CH-CH=CH-		H	methyl	oil
47	0	methyl	Br	n-propyl	methyl	69

48	0	methyl	Cl	H	methyl	98
49	0	methyl	I	H	methyl	102
50	0	methyl	Br	methyl	methyl	97
51	0	methyl	NO ₂	H	methyl	77
52	0	methyl	methoxy	H	methyl	135-137
53	0	methyl	Br	H	ethyl	
54	0	methyl	Br	n-butyl	methyl	
55	0	methyl	Br	phenyl	methyl	oil
56	0	methyl	Br	3-methyl-phenyl	methyl	oil
57	0	methyl	Br	2-methyl-propyl	methyl	90
58	0	methyl	Cl	H	ethyl	131-131
59	0	methyl	Cl	H	n-propyl	79
60	0	F	H	H	methyl	oil
61	0	Br	H	H	methyl	oil
62	0	methyl	methyl	H	methyl	63-64

35 Example 63Preparation of 5-bromo-6,6'-dimethyl-2,2'3',4'-tetramethoxy-benzophenone63A 6-methyl-2-methoxybenzoic acid

40 [0086] A mixture of ethyl 6-methyl-2-methoxybenzoate (642.0 g, 3.3 mol), water (2.51), ethanol (4.0 l) and sodium hydroxide (270 g, 6.6 mol) is heated under reflux with stirring for 20 hours. Subsequently, ethanol is distilled off and the reaction mixture is diluted with water, acidified with concentrated hydrochloric acid. The solid material is collected by vacuum filtration, washed with water and dried yielding off-white crystals, 460.0 g (83.9 %).

63B 5-Bromo-6-methyl-2-methoxybenzoic acid

45 [0087] A mixture of bromine (102 ml, 2.0 mol) and acetic acid (225 ml) is added to a mixture of 63A (304.0 g, 1.8 mol), sodium acetate (164.0 g, 2.0 mol) and acetic acid (3.0 l) at a temperature of 10 to 15 °C. The reaction mixture is stirred at room temperature for 16 hours. The solid material is collected by vacuum filtration, washed with water and dried yielding off-white crystals 321.0 g (72.6 %), mp 81-83 °C.

63C 5-bromo-6,6'-dimethyl-2,2'3',4'-tetramethoxy-benzophenone

55 [0088] 63B (240 g, 1.0 mol) is reacted with 3,4,5-trimethoxytoluene (182 g; 1.0 mol) in the presence of P₂O₅ (1.0 kg) and dichloromethane as described in Example 6 yielding white crystals, 220 g (54 %), mp 89-91 °C.

[0089] The sequence of the saponification (with sodium hydroxide) and bromination steps described above in Examples 63A and 63B, respectively, are reversed in Examples 6B and 6A, above.

[0090] The bromination described in Examples 63B and 6A is with the free acid and ester, respectively. The reaction conditions also differ, such as Example 63B which uses a polar protic solvent (acetic acid) and a buffering salt (sodium acetate). Example 6A uses a non-polar aprotic solvent (tetrachloromethane).

5 **Example 64**

Preparation of 6,6'-dimethyl-2,2'3',4'-tetramethoxy-5-trifluoromethyl-benzophenone

[0091] A mixture of 63C (6.15 g, 0.015 mol), sodium trifluoroacetate (9.52 g, 0.07 mol), copper-(I) iodide (13.37 g, 0.07 mol) and 1-N-methyl-2-pyrrolidinone (100 ml) is heated to 160 °C for 118 hours. The reaction mixture is cooled to room temperature and diluted with ether and water. The organic phase is separated and concentrated and the residue is purified by column chromatography (petrol ether : ethyl acetate, 95:5 v/v) yielding the pure product as white crystals, 1.3 g, (22 %) having a melting point of 95-96 °C.

15 **Example 65**

Preparation of 6'-methyl-2,2',3',4'-tetramethoxy-6-trifluoromethyl-benzophenone

65A 2-methoxy-6-trifluoromethylbenzoic acid

[0092] A mixture of potassium methylate in methanol (15 ml, 15 %) and 2-fluoro-6-trifluoromethylbenzoic acid (2.08 g, 0.01 mol) is heated to reflux with stirring for 40 hours. Upon cooling to room temperature water is added. The reaction mixture is filtered and the filtrate is acidified with concentrated hydrochloric acid. The solid material is collected by filtration, washed with water and dried yielding white crystals, 1.4 g (64 %), mp 130-131°C.

25 65B 6'-methyl-2,2'3',4'-tetramethoxy-6-trifluoromethyl-benzophenone

[0093] 65A (1.38 g, 0.00625 mol) is reacted with 3,4,5-trimethoxytoluene (1.14 g; 0.00625 mol) in the presence of P_2O_5 (5.0 g) and dichloromethane (100 ml) as described in Example 6 yielding white crystals, 1.68 g (69.8 %), mp 69-70 °C.

Example 66

Preparation of 5-chloro-6,6'-dimethyl-3'-ethoxy-2,2',4'-trimethoxybenzophenone

66A 5-chloro-6,6'-dimethyl-3'-propionyloxy-2,2',4'-trimethoxybenzophenone

[0094] 5-Chloro-2-methoxy-6-methylbenzoic acid (6.00 g, 0.03 mol) is reacted with 3,5-dimethoxy-4-propionyloxy-toluene (6.73 g; 0.03 mol) in the presence of P_2O_5 (18.0 g) and dichloromethane (200 ml) as described in Example 6 yielding white crystals, 6.65 g (56.4 %), mp 150-151 °C.

66B 5-chloro-6,6'-dimethyl-3'-hydroxy-2,2',4'-trimethoxybenzophenone

[0095] A mixture of 66A (6.65 g), potassium carbonate (7.0 g), water (50 ml) and methanol (100 ml) is heated to reflux for 3 hours. Upon cooling to room temperature water is added. The reaction mixture is filtered and the filtrate is acidified with concentrated hydrochloric acid. The solid material is collected and used without further purification.

66C 5-chloro-6,6'-dimethyl-3'-ethoxy-2,2',4'-trimethoxybenzophenone

[0096] A mixture of 66B (1.15 g), potassium carbonate (1.0 g), diethylsulfate (0.54 g) and acetonitrile (20 ml) is heated to reflux for 3 hours. Upon cooling to room temperature water is added. The solid material is collected and the residue is re-crystallized from methanol yielding white crystals, 1.07 g (85.6 %), mp 131-132 °C.

Example 67

Preparation of 6,6'-dimethyl-5-fluoro-2,2',3',4'-tetramethoxybenzophenone 67A 2-tert-butyl-4-fluoromethylphenol

[0097] A mixture of 4-fluoro-3-methylphenol (12.60 g, 0.1 mol), tert-butylchloride (25 ml) and $FeCl_3$ is heated to reflux

for 16 hours. The excess of tert-butylchloride is distilled off and the residue is purified by column chromatography (petrol ether : ethyl acetate, 95:5 v/v) yielding the pure product as a yellow oil, 15.3 g (84 %).

67B 6-bromo-2-tert-butyl-4-fluoro-5-methylphenol

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[0098] A mixture of 67A (0.91 g), tetrachloromethane (20 ml), N-bromosuccinimide (0.89 g) and AlCl_3 is stirred at ambient temperature for 3 days. The reaction mixture is filtered and concentrated. The obtained yellow oil (1.25 g) is used without further purification.

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67C 2-bromo-4-fluoro-3-methylphenol

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[0099] A mixture of 67B (18.5 g), benzene (120 ml) and AlCl_3 (6.5 g) is heated to reflux for 5 hours. The reaction mixture is cooled to room temperature and diluted with ethyl acetate and water. The organic phase is separated and concentrated. The residue is purified by column chromatography (petrol ether: ethyl acetate, 95:5 v/v) yielding the pure product as white crystals 9.0 g, (62 %).

67D 2-bromo-6-fluoro-3-methoxytoluene

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[0100] A mixture of 67C (9.0 g), potassium carbonate (6 g), dimethylsulfate (6 g) and acetonitrile (150 ml) is heated to reflux for 1 hour. The reaction mixture is cooled to room temperature, diluted with water and extracted with diethyl ether twice. The organic phase is separated and concentrated and the residue (9.0 g) is used without further purification.

67E 3-fluoro-6-methoxy-2-methylbenzoic acid

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[0101] A solution of n-butyllithium in hexane (18.1 ml, 2.5 mol/l) is slowly added to a mixture of 67D (9.0 g) and tetrahydrofuran (90 ml) at -78°C. The resulting reaction mixture is saturated with carbon dioxide at -78°C and allowed to warm up to ambient temperature. The reaction mixture is poured into water, acidified and extracted with diethylether twice. The organic phase is extracted with an aqueous solution of sodium hydroxide (5 %). The aqueous extract is acidified and extracted with diethylether twice. The organic phase is dried and concentrated. The residue (3.9 g) is used without further purification.

67F 6,6'-dimethyl-5-fluoro-2,2'3',4'-tetramethoxybenzophenone

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[0102] 67E (3.9 g, 0.021 mol) is reacted with 3,4,5-trimethoxytoluene (3.85 g; 0.021 mol) in the presence of P_2O_5 (12.0 g) and dichloromethane (150 ml) as described in Example 6 yielding white crystals, 0.34 g, mp 58°C.

Biological Investigations

A Foliar Systemicity

40

[0103] Wheat Powdery Mildew (WPM):

HOST: Wheat (*Triticum aestivum* L.) variety Kanzler

PATHOGEN: Blumeria (*Erysiphe*) graminis DC. f.sp. *tritici* E. Marchal

45

TEST PROCEDURE:

[0104]

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1. Wheat seed (8/pot) is planted in 8 cm diameter plastic pots in the greenhouse.
2. When the primary leaf is fully expanded, the plants are cut back to four in each pot of which two are marked with a permanent marker 5 cm below the leaf tip on the upper leaf surface. Thus there are two band-treated and two untreated plants in each pot.
3. A pipette is used to apply 5 μl of the formulated compound in a band on the lower leaf surface opposite the mark. The application band should cover the whole leaf width. After application, the plants are not moved until the bands are dry (half an hour or so later).
4. After treated plants have dried, they are moved to the greenhouse and kept there for 2 days to allow for movement of the compounds. The plants are maintained with bottom watering.

5. Two days after application, the plants are inoculated by dusting them with powdery mildew conidia in the green-house. Evaluations are typically made 7 - 8 days after inoculation.

Evaluation

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[0105] Three types of compound movement are assessed by evaluating disease in three areas of each band-treated leaf.

10

[0106] Translaminar movement: The percent disease area is assessed for the translaminar band area (marked area of the upper leaf surface directly opposite where band was applied on the lower leaf surface; width of band approximately 5 mm). Translaminar disease control is then calculated using the following formula:

$$\% \text{ disease control} = 100 - \frac{\% \text{ disease in treated plants}}{\% \text{ disease in untreated plants}} \times 100$$

15

[0107] Distal movement and proximal movement: The distal and proximal disease-free zones on the upper leaf surface are measured in mm. The distal direction is from the band toward the leaf apex and the proximal direction is from the band toward the leaf base. The percent of the disease-free zone relative to the entire distance between the band and leaf apex or base is calculated. If disease is noticeably lighter in the distal or proximal area this is also noted.

20

FORMULATION AND CONTROLS:

[0108]

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1. The compounds are formulated in a solvent/surfactant system consisting of 5% acetone and 0.05% Tween 20 in deionized water. Formulated compounds are prepared using deionized water. Compounds are typically tested at 400 ppm.

2. Two kinds of controls are included:

30

- Plants band-treated with the solvent/surfactant solution and inoculated (Solvent Blank).
- Untreated plants which are inoculated (Inoculated Control).

[0109] The results of this evaluation are shown in Table III:

TABLE III

35

Foliar Systemicity

40

Example No.	Proximal Movement (mm from band)	Distal Movement (mm from band)	Translaminar Activity (%)
1	10	50	100
10	6	46	100
Standard ¹	4	5	100
Standard ²	8	28	100

45

The following compounds, which are known from EP 0 727 141 have been used as standards:

Standard¹ 2,6-dichloro-6'-methyl-2',3',4'-trimethoxybenzophenone
 Standard² 2',3',4'-trimethoxy-2,6,6'-trimethylbenzophenone

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B-1 Comparison of the fungicidal activity of the 2-methoxy-benzophenones to a 2,6-dichloro- and 2,6-dimethyl-benzophenone

Test diseases:

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[0110]

(a) Wheat Powdery Mildew (WPM):

HOST: Wheat (*Triticum aestivum* L.) variety Kanzler
PATHOGEN: *Blumeria (Erysiphe) graminis* DC. f.sp. *tritici* E. Marchal

5 (b) Barley Powdery Mildew (BPM):

HOST: Barley (*Hordeum vulgare* L.) variety Golden Promise
PATHOGEN: *Blumeria (Erysiphe) graminis* DC. f.sp. *hordei* E. Marchal

10 TEST PROCEDURE:

[0111] This test is a zero day protectant test for control of wheat and barley powdery mildews.

1. Wheat or barley seed (approximately 8-10/pot) is planted in 6 cm diameter plastic pots and maintained in the greenhouse.
- 15 2. When the primary leaf is fully expanded, formulated test compounds are sprayed with a single nozzle overhead track sprayer at a rate of 200 l/ha. Plants are then allowed to air-dry.
3. Inoculation follows about three hours after compound application. Plants are set up on greenhouse benches with bottom watering mats and inoculated by dusting them with conidia from powdery mildew infected plants (stock cultures at an age of 10-14 days).
- 20 4. Disease on the primary leaf as percent leaf area with disease symptoms/signs is evaluated about 7 days after inoculation. The tips and bases of the leaves are excluded from the evaluation. Percent disease control is then calculated by the following formula:

$$25 \quad \% \text{ disease control} = 100 - \frac{\% \text{ infection in treated plants}}{\% \text{ infection in untreated plants}} \times 100$$

FORMULATION, REFERENCE COMPOUNDS AND CONTROLS:

30 [0112]

1. Technical compounds are formulated in a solvent/surfactant system consisting of 5% acetone and 0.05% Tween 20 in deionized water. Compounds are dissolved in acetone prior to addition of the water; the Tween 20 can be added through either the acetone or the water. Dilutions are made using the solvent/surfactant system. Test compounds are typically tested over a range of concentrations covering several orders of magnitude and then ED values are calculated for comparison of compounds.
- 35 Formulated compounds are prepared using deionized water.
2. Two kinds of controls are included:

40 Plants treated with the solvent/surfactant solution and inoculated (Solvent Blank).
Untreated plants which are inoculated (Inoculated Control).

[0113] The results of this evaluation are shown in Table IV:

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50

55

Table IV

Fungicidal activity of 2-methoxybenzophenones (ED90 values)										
Comparison of the fungicidal activity of the 2-methoxy-benzophenones to a 2,6-dichloro- and 2,6-dimethyl benzophenone										
Results are from 0 day protectant tests in which all analogs were tested side-by-side.										
Disease			ED90 (ppm)							
	Standard ¹	Standard ²	1	4	6	7	10	48	Quinoxifen	
WPM	28	20	4	5	0.1	7	7	0.1	12	
BPM	24	8	6	7	0.9	6	8	<0.1	26	

[0114] Compounds applied as technical material formulated in 0.5% acetone 0.05% Tween 20 water

B-2 Comparison of the curative and residual fungicidal activity of the 2-methoxy-benzophenones to a 2,6-dichloro- and 2,6-dimethylbenzophenone

Test diseases:

[0115]

(a) Wheat Powdery Mildew (WPM):

HOST: Wheat (*Triticum aestivum* L.) variety Kanzler

PATHOGEN: *Blumeria (Erysiphe) graminis* DC. f.sp. *tritici* E. Marchal

(b) Cucumber Powdery Mildew (QPM):

HOST: Cucumber (*Cucumis sativus* L.) variety Bush pickle

PATHOGEN: *Erysiphe cichoracearum* DC

TEST PROCEDURE:

[0116] This test procedure is for curative and residual control of powdery mildew diseases.

1. Wheat seed (approximately 8-10/pot) or cucumber seed (1 seed / pot) is planted in 6 cm diameter plastic pots and maintained in the greenhouse.

2. When the primary leaf (wheat) or the cotyledons (cucumber) is/are fully expanded, formulated test compounds are sprayed with a single nozzle overhead track sprayer at a rate of 200 l/ha. Plants are then allowed to air-dry.

3. Inoculation precedes treatment by 2 or 3 days in the case of curative evaluations and follows treatment by 3 or 4 days in case of residual evaluations. For inoculation, plants are set up on greenhouse benches with bottom watering mats and inoculated by dusting them with conidia from powdery mildew infected plants (stock cultures at an age of 10-14 days). Between inoculation and treatment for curative evaluations and between treatment and inoculation for residual evaluations, plants are maintained in the greenhouse with bottom watering.

4. Disease on the primary leaf (wheat) or on the cotyledons (cucumber) as percent leaf area with disease symptoms/signs is evaluated about 7 days after inoculation. In the case of wheat, the tips and bases of the leaves are excluded from the evaluation. Percent disease control is then calculated by the following formula:

$$\% \text{ disease control} = 100 - \frac{\% \text{ infection in treated plants}}{\% \text{ infection in untreated plants}} \times 100$$

FORMULATION, REFERENCE COMPOUNDS AND CONTROLS:

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[0117]

10

1. Technical compounds are formulated in a solvent/surfactant system consisting of 5% acetone and 0.05% Tween 20 in deionized water. Compounds are dissolved in acetone prior to addition of the water; the Tween 20 can be added through either the acetone or the water. Dilutions are made using the solvent/surfactant system. Formulated compounds are prepared using deionized water.

2. Two kinds of controls are included:

15

Plants treated with the solvent/surfactant solution and inoculated (Solvent Blank).

Untreated plants which are inoculated (Inoculated Control).

[0118] The results of this evaluation are shown in Tables V and VI:

Table V

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Curative and Residual Fungicidal activity of 2-methoxybenzophenones compared with standard compounds							
Disease Test	Rate (ppm)	Disease control (% efficacy)					
		Standard 1		Standard 2		2-methoxy benzophenone Example	
	Test	A	B	A / B	A / B	B	C
WPM 2 da C	1250	79	92	88/85	98/87	61	97
	125	60	74	79 / 65	95 / 89	71	93
	12,5	31	55	69 / 52	90/61	55	73
WPM 3 da R	1250	100	100	100/100	100/100	100	100
	125	83	81	100 / 100	100 / 100	100	100
	12,5	75	70	90/100	100 / 100	99	99
QPM 3 da R	1250	100	89	100/100	100 / 100	100	100
	125	0	6	5 / 61	92 / 92	97	89
	12,5	0	14	2/3	2/35	8	2

2 da C = 2 day curative inoculation 2 days BEFORE application 3 da R = 3 day residual inoculation 3 days AFTER application

Table VI

45

Curative and Residual Fungicidal activity of 2-methoxy-benzophenones against wheat powdery mildew						
Test	Rate (ppm)	Disease control (% efficacy) 2-methoxy-benzophenone Example				
		6	58	59	64	65
WPM 3 da C	125	64	66	69	72	58
	25	60	58	63	60	48
	5	43	48	48	48	23
	1	27	29	28	37	23
	0.2	17	17	13	10	10

Table VI (continued)

Curative and Residual Fungicidal activity of 2-methoxy-benzophenones against wheat powdery mildew						
Test	Rate (ppm)	Disease control (% efficacy) 2-methoxy-benzophenone Example				
WPM 4 da R	125	100	100	100	100	100
	25	100	100	100	100	99
	5	100	100	100	99	93
	1	90	95	95	89	50
	0.2	58	68	85	51	25

3 da C = 3 day curative Inoculation 3 days BEFORE application 4 da R = 4 day residual Inoculation 4 days AFTER application

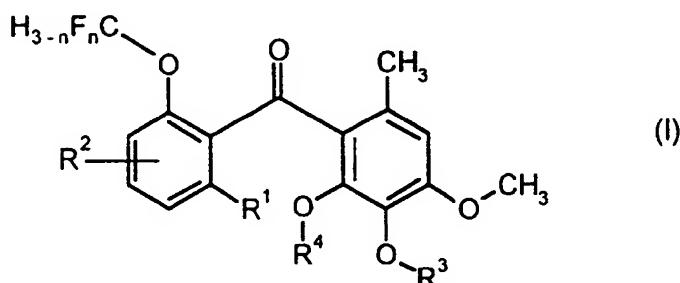
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Claims

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1. A compound of formula I:

25



30

wherein

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R¹ represents a halogen atom or an alkyl group or haloalkyl group;
 R² represents a hydrogen or halogen atom, or an alkyl, haloalkyl, alkoxy or nitro group; or R¹ and R² together represent a group of formula -CH=CH-CH=CH-;
 R³ and R⁴ each independently represent an optionally substituted alkyl; and
 n is an integer from 0 to 3.

40

2. A compound as claimed in claim 1, wherein

45

R¹ represents a chloro atom or a methyl or trifluoromethyl group;
 R² represents a hydrogen, chloro or bromo atom or a trifluoromethyl group;
 R³ represents a C₁₋₅ alkyl group;
 R⁴ represents a C₁₋₅ alkyl group or benzyl group, being optionally substituted by at least one halogen atom, or C₁₋₄ alkyl or alkoxy group; and
 n is 0.

50

3. A compound as claimed in claim 1, wherein at least one of the radicals R¹ and R² represents a trifluoromethyl group.

55

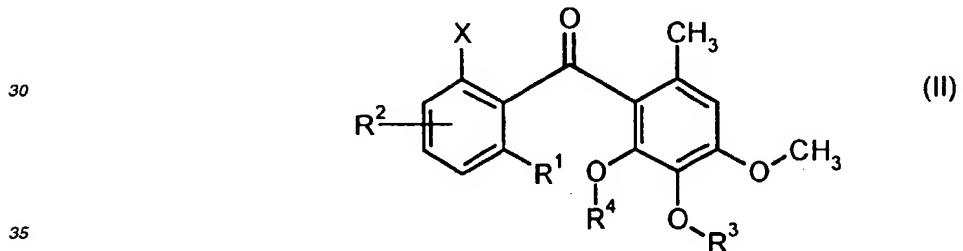
4. A compound as claimed in claim 1 being selected from the group consisting of

6,6'dimethyl-2,2',3',4'-tetramethoxy-benzophenone,
 6,6'-dimethyl-3'-pentoxy-2,2',4'-trimethoxy-benzophenone,
 5-bromo-6,6'-dimethyl-2,2',3',4'-tetramethoxy-benzophenone,
 5-chloro-6,6'-dimethyl-2,2',3',4'-tetramethoxy-benzophenone,
 5-iodo-6,6'-dimethyl-2,2',3',4'-tetramethoxy-benzophenone,

6-chloro-6'-methyl-2,2',3',4'-tetramethoxy-benzophenone,
 5-bromo-6-chloro-6'-methyl-2,2',3',4'-tetramethoxy-benzophenone,
 6-chloro-5,6'-dimethyl-2,2',3',4'-tetramethoxy-benzophenone,
 2'-butoxy-6-chloro-6'-methyl-2,2',3',4'-trimethoxy-benzophenone,
 5 2'-butoxy-6-chloro-5,6'-dimethyl-2,2',3',4'-trimethoxy-benzophenone
 6-chloro-2'-(2-fluorobenzyl)oxy-6'-methyl-2,2',3',4'-trimethoxy-benzophenone,
 6-chloro-2'-(4-fluorobenzyl)oxy-6'-methyl-2,2',3',4'-trimethoxy-benzophenone,
 5-bromo-6,6'-dimethyl-3'-pentyloxy-2,2',4'-trimethoxy-benzophenone
 6-chloro-6'-methyl-2'-pentyloxy-2,2',3',4'-trimethoxy-benzophenone,
 10 6-chloro-2'-(3-methylbutyloxy)-6'-methyl-2,2',3',4'-trimethoxy-benzophenone,
 2'-benzyloxy-6-chloro-6'-methyl-2,2',3',4'-trimethoxy-benzophenone,
 6-chloro-2'-(3-methylbenzyloxy)-6'-methyl-2,2',3',4'-trimethoxy-benzophenone,
 6-chloro-2'-(4-methylbenzyloxy)-6'-methyl-2,2',3',4'-trimethoxy-benzophenone,
 15 6-chloro-2-difluoromethoxy-6'-methyl-2',3',4'-trimethoxy-benzophenone,
 1-(6'-methyl-2',3',4'-trimethoxybenzoyl)-2-methoxynaphthalene,
 1-(6'-methyl-2',3',4'-Anmethoxybenzoyl)-2-difluoromethoxynaphthalene,
 5-chloro-6,6'-dimethyl-3'-ethoxy-2,2',4'-trimethoxy-benzophenone,
 5-chloro-6,6'-dimethyl-3'-n-propyloxy-2,2',4'-trimethoxy-benzophenone,
 6,6'-dimethyl-2,2',3',4'-tetramethoxy-5-trifluoromethyl-benzophenone,
 20 6'-methyl-2,2',3',4'-tetramethoxy-6-trifluoromethyl-benzophenone,
 5-bromo-6'-methyl-2,2',3',4'-tetramethoxy-6-trifluoromethyl-benzophenone.

5. A process for the preparation of the compound of claim 1, wherein n is 0, which comprises reacting a compound of formula II:

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wherein R¹, R², R³ and R⁴ are as described in claim 1, and X represents a fluoro or chloro atom, with an alkali methanolate.

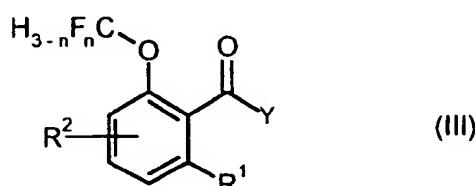
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6. A process for the preparation of the compound of claim 1, which comprises

(a) reacting a compound of formula III:

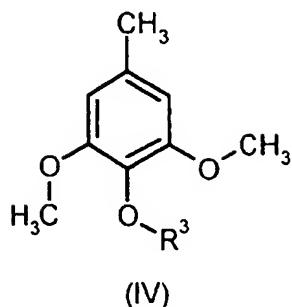
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50



wherein R¹, R² and n are as described in claim 1, and Y represents a leaving group or a hydroxy group, with a compound of formula IV:

55



15 wherein R³ is as described in claim 1, in the presence of a Lewis acid or a dehydrating agent; and
 (b) optionally treating the product obtained with a compound of formula V:

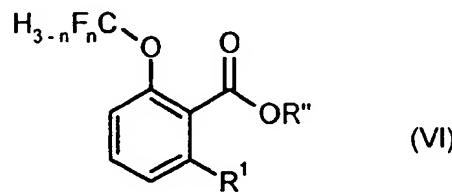


20 wherein R⁴ represents an optionally substituted alkyl or aryl group with the proviso that the alkyl group is not methyl, and Met denotes an alkali metal atom.

25

7. A process for the preparation of a compound of formula I, wherein one of the radicals R¹ and R² represents a trifluoromethyl group, which comprises treating a compound of formula I, wherein one of the radicals radicals R¹ and R² represents a halogen atom with trifluoroacetic acid in the presence of a copper salt at elevated temperatures.
8. A process for the preparation of a formula III compound of claim 6, wherein R² represents a halogen atom, which comprises

30 (a) reacting a compound of formula VI:

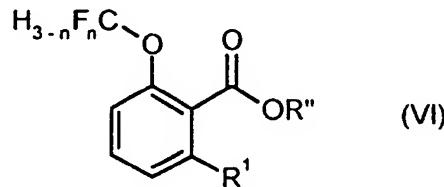


40 wherein R¹ and n are as described in claim 1, and R'' represents a hydrogen atom or an alkyl group, with a halogenating agent.

45

9. A process for the preparation of a formula III compound of claim 6, wherein R² represents a halogen atom, which comprises

(a) reacting a compound of formula VI:



wherein R¹ and n are as described in claim 1, and R^a represents an alkyl group, with a halogenating agent; (b) optionally hydrolyzing the resulting halogenated alkyl benzoate; and (c) optionally treating the resulting halogenated alkyl benzoic acid with thionyl chloride.

5 10. A composition which comprises a fungicidally effective amount of at least one compound of Claim 1, and a carrier.

11. A method of combating fungus or a fungal plant disease at a locus, which comprises treating the locus with a compound of claim 1.

10 12. A method of combating fungus or a fungal plant disease at a locus, which comprises treating the locus with the composition of claim 10.

13. The method of claim 11 wherein the fungal plant disease is ascomycetes.

15 14. The method of claim 12 wherein the fungal plant disease is ascomycetes.

15. The method according to claim 11 wherein the fungus is a member of the subgroup Erysiphales

20 16. The method of Claim 15 wherein the fungus is selected from *Blumeria (Erysiphe) graminis*, *Erysiphe cichoracearum*, *Podosphaera leucotricha* or *Uncinula necator*.

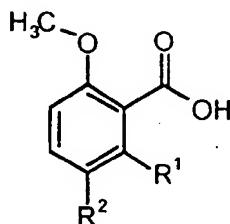
17. The method according to claim 12 wherein the fungus is a member of the subgroup *Erysiphales*.

25 18. The method of claim 17 wherein the fungus is selected from *Blumeria (Erysiphe) graminis*, *Erysiphe cichoracearum*, *Podosphaera leucotricha* or *Uncinula necator*.

30 19. A process for the preparation of a compound of formula III, wherein R² represents a bromine atom or trifluoromethyl group, according to claim 7, which comprises reacting a compound of formula VI, wherein R¹ is an alkyl group; n is 0; and R^a represents a hydrogen atom, with bromine and optionally treating the resulting bromo compound with trifluoroacetate in the presence of a copper salt at elevated temperatures.

20 20. A process for the preparation of a compound of formula III according to claim 19, wherein the reaction between the compound of formula VI and bromine is carried out in the presence of a polar protic solvent.

35 21. A compound having the formula



wherein R¹ represents an alkyl group and R² represents a bromo atom or a trifluoromethyl group.

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European Patent
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EUROPEAN SEARCH REPORT

Application Number
EP 98 30 6583

DOCUMENTS CONSIDERED TO BE RELEVANT			CLASSIFICATION OF THE APPLICATION (Int.Cl.6)						
Category	Citation of document with indication, where appropriate, of relevant passages	Relevant to claim							
D, X	EP 0 727 141 A (AMERICAN CYANAMID CO) 21 August 1996 * the whole document * -----	1-3, 6, 10-18	C07C49/84 C07C205/45 C07C65/21 A01N35/04						
X	ISLAM M R ET AL: "A new synthetic route to 4-bromo-7-methoxyphthalide and 3-hydroxy-7-methoxyphthalide from 2-methoxy-6-methylbenzoate: side-chain and nuclear bromination" J. CHEM. RES., SYNOP. (JRPSDC, 03082342);91; (2); PP.29, XP002086460 Manchester Univ.; Chem. Dep.; Manchester; M13 9PL; UK (GB) * page 29 * -----	8	C07C45/46 C07C45/70 C07C45/68 C07C51/363 C07C67/307 C07C51/60 C07C51/347 C07C67/343 C07C69/92						
A		21							
			TECHNICAL FIELDS SEARCHED (Int.Cl.6)						
			C07C A01N						
<p>The present search report has been drawn up for all claims</p> <table border="1" style="width: 100%; border-collapse: collapse;"> <tr> <td style="width: 33%;">Place of search</td> <td style="width: 33%;">Date of completion of the search</td> <td style="width: 34%;">Examiner</td> </tr> <tr> <td>THE HAGUE</td> <td>2 December 1998</td> <td>Bonneville, E</td> </tr> </table>				Place of search	Date of completion of the search	Examiner	THE HAGUE	2 December 1998	Bonneville, E
Place of search	Date of completion of the search	Examiner							
THE HAGUE	2 December 1998	Bonneville, E							
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ANNEX TO THE EUROPEAN SEARCH REPORT
ON EUROPEAN PATENT APPLICATION NO.

EP 98 30 6583

This annex lists the patent family members relating to the patent documents cited in the above-mentioned European search report. The members are as contained in the European Patent Office EDP file on. The European Patent Office is in no way liable for these particulars which are merely given for the purpose of information.

02-12-1998

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